Attorney Docket No. 89188.0022

Amdt. Dated October 23, 2008

Customer No.: 26021

Reply to Advisory Action of August 22, 2008 and Final Office Action of April 24,

2008

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in

the application:

Listing of Claims:

1-31. (Canceled)

32. (Currently amended) A method for inhibiting the expression of a target

gene through an intracellular-RNA interference a post-transcriptional gene

silencing mechanism in a cell or organism that expresses the targeted gene,

comprising the steps of:

a) providing a composition comprising an mRNA-cDNA hybrid duplex

prior to contacting said cell or said organism, wherein the mRNA-cDNA hybrid

duplex is capable of used by the cell or organism as a template to generate small

gene-silencing effectors inhibiting the expression of said targeted gene in said cell or

organism, wherein said expression is inhibited through said intracellular RNA

interference post-transcriptional gene silencing mechanism; and

b) contacting said cell or said organism with said composition under

conditions such that the expression of said gene in said cell or said organism is

inhibited,

wherein the mRNA is a ribonucleic acid sequence in the sense orientation of

said targeted gene and the cDNA is a deoxyribonucleic acid sequence in the anti-

sense orientation of said targeted gene, and wherein the mRNA-cDNA hybrid

duplex forms between said mRNA and said cDNA in a complementary region

containing more than 500 base pairs.

33. (Canceled)

Page 2 of 13

Attorney Docket No. 89188.0022

Amdt. Dated October 23, 2008

Customer No.: 26021

Reply to Advisory Action of August 22, 2008 and Final Office Action of April 24,

2008

34. (Previously presented) The method of Claim 32, wherein said cell or

said organism expresses said targeted gene in vivo, wherein said cell or said

organism is a cell, tissue, or organ selected from the group consisting of cancerous

cells, liver, and skin.

35. (Currently amended) The method of Claim 32, wherein said targeted

gene comprises a gene selected from the group consisting of sonic hedgehog (Shh),

beta-catenin, and bel-2 gene.

36. (Currently amended) The method of Claim 32, wherein said mRNA-

cDNA hybrid duplex inhibits the expression of said targeted gene, wherein said

targeted gene comprises a region selected-from-the sonic hedgehog sequence

amplified by SEQ ID NOs: 12, 13, 14, and 15, a \(\theta\)-catenin sequence encoding its

amino acid domain from position 306 to 644, and a bel-2 sequence amplified by SEQ

ID NOs: 8 and 9.

37. (Canceled)

38. (Previously presented) The method of Claim 32, wherein said cell is a

prokaryotic cell.

39. (Canceled)

40. (Previously presented) The method of Claim 38, wherein said cell is a

bacterial cell.

41. (Previously presented) The method of Claim 32, wherein said cell or

said organism is a cell or organism of an eukaryote.

Page 3 of 13

Attorney Docket No. 89188.0022

Amdt. Dated October 23, 2008

Customer No.: 26021

Reply to Advisory Action of August 22, 2008 and Final Office Action of April 24,

2008

42. (Original) The method of Claim 41, wherein said eukaryote is a

vertebrate.

43. (Original) The method of Claim 41, wherein said eukaryote is a mouse.

44. (Original) The method of Claim 41, wherein said eukaryote is

chimpanzee.

45. (Original) The method of Claim 41, wherein said eukaryote is a human

being.

46-54. (Canceled)

55. (Previously presented) The method of Claim 32, wherein the

composition consists of an mRNA-cDNA hybrid duplex capable of inhibiting the

expression of said targeted gene, wherein the mRNA is a ribonucleic acid sequence

in the sense orientation of said targeted gene and the cDNA is a deoxyribonucleic

acid sequence in the anti-sense orientation of said targeted gene, wherein the

mRNA-cDNA hybrid duplex forms between said mRNA and said cDNA in a

complementary region containing more than 500 base pairs.

56-57. (Canceled)

58. (Previously presented) The method of Claim 55, wherein said targeted

gene is expressed in a cell or organism in vivo, wherein said cell or said organism is

a cell, tissue, or organ selected from the group consisting of cancerous cells, liver,

and skin.

Attorney Docket No. 89188.0022

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59. (Currently amended) The method of Claim 55, wherein said targeted

gene comprises a gene-selected from the group consisting of sonic hedgehog (Shh),

beta-catenin, and bel-2 gene.

60. (Currently amended) The method of Claim 55, wherein said mRNA-

cDNA hybrid duplex inhibits the expression of said targeted gene, wherein said

targeted gene comprises a region selected from the sonic hedgehog sequence

amplified by SEQ ID NOs: 12, 13, 14, and 15, a \(\theta\)-catenin sequence encoding its

amino acid domain from position 306 to 644, and a bel 2 sequence amplified by SEQ

ID NOs: 8 and 9.

61. (Previously presented) The method of Claim 55, wherein said cell is a

prokaryotic cell.

62. (Canceled)

63. (Previously presented) The method of Claim 61, wherein said cell is a

bacterial cell.

64. (Previously presented) The method of Claim 55, wherein said cell or

said organism is a cell or organism of an eukaryote.

65. (Previously presented) The method of Claim 64, wherein said

eukaryote is a vertebrate.

66. (Previously presented) The method of Claim 64, wherein said

eukaryote is a mouse.

Page 5 of 13

Attorney Docket No. 89188.0022

Amdt. Dated October 23, 2008

Customer No.: 26021

Reply to Advisory Action of August 22, 2008 and Final Office Action of April 24,

2008

67. (Previously presented) The method of Claim 64, wherein said eukaryote is chimpanzee.

- 68. (Previously presented) The method of Claim 64, wherein said eukaryote is a human being.
- 69. (Currently amended) A method for inhibiting the expression of a target gene through an intracellular RNA interference a post-transcriptional gene silencing mechanism in a cell or organism that expresses the targeted gene, comprising the steps of:
- a) providing a composition comprising an mRNA-cDNA hybrid duplex prior to contacting said cell or said organism, wherein the mRNA-cDNA hybrid duplex is eapable of used by the cell or organism as a template to generate small gene-silencing effectors inhibiting the expression of said targeted gene in said cell or said organism, wherein said expression is inhibited through said intracellular RNA interference post-transcriptional gene silencing mechanism; and
- b) contacting said cell with said composition under conditions such that the expression of said gene in said cell is inhibited,

wherein the mRNA is a ribonucleic acid sequence in the sense orientation of said targeted gene and the cDNA is a deoxyribonucleic acid sequence in the antisense orientation of said targeted gene, and wherein the mRNA is a full-length transcript of said targeted gene larger than 500 base pairs.

70. (Previously presented) The method of Claim 69, wherein the mRNA is an unspliced mRNA transcript of the targeted gene.

Appl. No. 09/920,342

Amdt. Dated October 23, 2008

Reply to Advisory Action of August 22, 2008 and Final Office Action of April 24, 2008

71. (Previously presented) The method of Claim 69, wherein the mRNA is a spliced mRNA transcript of the targeted gene.